Page 1 of 2

Form PTO-1449 (MODIFIED) U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE

ATTY. DOCKET NO. 038602-1081 SERIAL NO.

09/769,360

INFORMATION DISCLOSURE CITATION

Peng C. TANG et al.

FILING DATE

APPLICANT

GROUP ART UNIT

(Use several sheets if necessary)

01/26/2001

1362 1624

U.S. PATENT I	DOCUMENTS
---------------	-----------

EX	MITALE	REF	DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE IF APPROPRIATE	
	120	<u> </u>	RE 36,256	07/99	Spada et al.	514	249		
9	IN 132 200	٢	5,217,999	06/93	Levitzki et al	514	357	문	
(3)	BC	9	5,302,606	04/94	Spada et al.	514	357	0 -	韶
1	PADEMARY B		5,330,992	07/94	Eissenstat et al.	514	312	N N	The state of the
	BC	pa-45	5,430,148	07/95	Webber et al	514	238	1 4 ER	血
	BC		5,534,518	07/96	Henrie et al	514	260	20	\leq
	BC		5,714,493	02/98	Myers et al	514	259	2001	
	BC		5,866,572	02/99	Barker et al	514	234.5	8	

FOREIGN PATENT DOCUMENTS

		REE DOCUMENT DATE COUNTRY	CLASS	SUB-	TRANSLATION			
	REF	NUMBER :	DATE	COUNTRY	CLASS	CLASS	YES	NO
BC		99/10325	04/99	WIPO/PCT				
BC		566 226 A1	10/93	European				
BC		91/15495	10/91	WIPO				
BC		92/21660	12/92	WIPO				
BC		92/20642	11/92	WIPO				
BC		94/03427	02/94	WIPO		17	紹	
BC		94/14808	07/94	WIPO		8 =	(C)	
BC		96/22976	08/96	WIPO		<u> Α</u> ω	W	
BC		96/40116	12/96	WIPO		200		

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc

- 41	Hynes et al., "Quinazolines as Inhibitors of Dihydrofolate Reductase", Journal of Medicinal Chemistry, Vol. 17
BC	No. 9, pages 1023-1025, (1974)
0.6	Chan et al., "Selective Inhibitors of Candida albicans Dihydrofolate Reductase: Activity and Selectivty of
Be	5-(Arylthiol)-2,4-diaminoquinazolines", J. of Med. Chem., vol. 38, No. 18, pages 3608-3616, (1995)
0.0	Bonner et al., "Structure and biological activity of human homologs of the raf/mil oncogene", Molecular and
- 13C	Cellular Biology, 5(6):1400-1407, (1985)

EXAMINER

Coleman

DATE CONSIDERED

August 16, 2002

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include any copy of this form with next communication to applicant.

Page 2 of 2

FOREIGN PATENT DOCUMENTS REF DOCUMENT DATE COUNTRY CLASS SUB-CLASS VES NO OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) Harfo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide dargeted against C-raf Kinase", Nature medicine, 2:668-675 (1998) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1986) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	Form PTO-1449 U.S. DEPARTMENT OF COMMERCE		ATTY. DOCKET NO.		SERIAL NO.					
INFORMATION DISCLOSURE CITATION Peng C. TANG et al. FILING DATE O1/26/2001 ONLOWERTS U.S. PATENT DOCUMENTS U.S. PATENT DOCUMENTS EXAMINER REF DOCUMENT NUMBER DATE NAME CLASS SUBCLASS PROPRIATE PENDITUDE PROPRIATE FOREIGN PATENT DOCUMENTS FOREIGN PATENT DOCUMENTS CLASS SUBCLASS PENDING PROPRIATE OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) Harfo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents 33(1):434-444, (1990) Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Whynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Whynes et al., "Cuinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Ckem. 32(3):569-575, (1988) Monia et al., "antifumor activity of a phosphorothioate antisense oligedeoxynucleotide largested against C-raf Kinase", Nature medicine, 2:688-675 (1996) Morrison et al. "Signal transduction from membrane to cytolasm: Growth factors and right page against C-raf Kinase", Nature medicine, 2:688-675 (1996) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and right page against C-raf Kinase", Nature medicine, 2:688-675 (1996) Robbins at al., "Regulation and properties of extracellular signal/oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	(MODIFIED)		PATENT AND TR	ADEMARK OFFICE			09/769,360			
U.S. PATENT DOCUMENTS EXAMINER REF DOCUMENT DATE NAME CLASS SUB- FILING DATE INTITUDE TOCUMENTS FOREIGN PATENT DOCUMENTS FOREIGN PATENT DOCUMENTS FOREIGN PATENT DOCUMENTS FOREIGN PATENT DOCUMENTS CLASS SUB- CLASS TRANSLATION CLASS TRANSLATION CLASS TRANSLATION CLASS TRANSLATION (THE DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) BC Hynes et al., "Turther studies on the synthesis of quinazolines from 2-fluorobenzonitriles", J. Heterocyclic Chem. 25:137-1363, (1991) BC Marsham et al., "Cuinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Class. 32(3):569-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity." Eric. Natl Acad. Sci. USA, 85:8855-8899 (1988) BC Robbins et al., "Regulation and properties of extracellular signal/dregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 288:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolineners", J. Med. Chem., 36(6):733-746, (1993)	(
U.S. PATENT DOCUMENTS EXAMINER REF DOCUMENT NUMBER DATE NAME CLASS SUB-CLASS PLOPROPRIATE FOREIGN PATENT DOCUMENTS REF DOCUMENT NUMBER DATE COUNTRY CLASS SUB-CLASS PLOPROPRIATE FOREIGN PATENT DOCUMENTS REF DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) Harfo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents 33(1):434-444, (1990) Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) BC Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic Chem. 25:1173-1177 (1986) BC Hynes et al., "Turther studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) BC Marsham et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotided and satilities", J. Med. Chem. 32(3):569-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and manufacture burned Concogene products increase Raf-1 phophorylation and associated protein kinase activity." Edic. Natl Acad. Sci. USA, 85:8855-8859 (1988) BC Robbins et al., "Regulation and properties of extracellular signal/oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 288:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinenes", J. Med. Chem., 36(6):733-746, (1993)	INFORMATION DISCLOSURE CITATION				Pe	ng C. TAN	G et al.			
U.S. PATENT DOCUMENTS EXAMINER INTITULE REF NUMBER DATE NAME CLASS SUBCLASS FILING DATE APPROPRIATE PENDICUMENT NUMBER DATE COUNTRY CLASS SUBCLASS FILING DATE COUNTRY CLASS SUBCLASS FOR NO THE NUMBER DATE COUNTRY CLASS SUBCLASS FOR NO THE POCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) BC Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic Chem. 25:1173-1177 (1988) BC Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1988) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-bodind Oncogene products increase Raf-1 phophorylation and associated protein kinase activity," Edge. Natl Acad. Sci. USA, 86:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal/dregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 286:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novet class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)				FILING DATE		GROUP AR	T UNIT			
FOREIGN PATENT DOCUMENTS OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) BC Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) BC Hynes et al., "Curinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 28:1357-1363, (1991) BC Morrison et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):559-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrage-bothed Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Edc. Natl Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novet class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)		(Use se	everal sheets if nece	ssary)	01/26/200)1		1362 1	<i>-</i> 24	
FOREIGN PATENT DOCUMENTS FOREIGN PATENT DOCUMENTS REF DOCUMENT DATE COUNTRY CLASS SUB-CLASS TRANSLATION YES NO OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) BC Hynes et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Heterocyclic chem. 25:1173-1177 (1988) BC Hynes et al., "Turcher studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) BC Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Cliem. 32(3):563-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide graphed against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Eds. Natl Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)				U.S. PATE	NT DOCUMENTS				,	
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):559-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-brand Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Exc. Natl Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signalOregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	1	REF	1	DATE	NAME	CLASS		300-		
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):559-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-brand Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Exc. Natl Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signalOregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	To the state of th	\								
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):559-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-brand Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Exc. Natl Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signalOregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	WH 1 2 2001 E	<u>y)</u>		FOREIGN PA	TENT DOCUMENTS		 	T-70.440		
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) Harlo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and hambagne-band Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Edc. Natl Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)		REF	1	DATE	COUNTRY	CLASS		ļ		
Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) Harlo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argisted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", 2nc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal Oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	THADEMAR							1.20		
Judah Folkman, "what is the evidence that tumors are angiogenesis dependent?", J. Natl. Cancer Institute 82(1):4-6, (1990) Harlo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argisted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", 2nc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal Oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)		· · · · · · · · · · · · · · · · · · ·	OTHER DOCU	MENTS (Including	Author Title Date Pert	inent Pages	Ftc.)	<u> </u>		
BC Harlo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) BC Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argeted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Eroc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) BC Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)								ncer Instit	ute	
Harlo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents BC Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide targeted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	BC				Tac turnors are angiogenes	sis dependent	- , J. Nau. Oa		ote	
Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem. 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide targeted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			Harlo & Lane, "Antibodies: A Laboratory manual, Cold Spring Harbor Laboratories, (1989) (Tables of Contents)							
BC 33(1):434-444, (1990) Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argued against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Eroc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	BC		The state of the s							
Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Clam. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argited against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			Harris et al., "Antifolate and antibacterial activities of 5-substituted 2,4-diaminoquinazolines", J. Med. Chem.							
Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem. 25:1173-1177 (1988) Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Clam. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argited against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	BC		33(1):434-444, (1990)							
Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem. 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide targeted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-band Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Eroc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal/oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			Hynes et al., "Direct synthesis of 2,4-Diaminoquinazolines from 2-fluorobenzonitriles", J. Heterocyclic chem.							
BC 28:1357-1363, (1991) Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide targeted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Enc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal Oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			25:1173-1177 (198	8)						
Marsham et al., "Quinazoline Antifolate Thymidylate Synthase Inhibitors", J. Med. Chem. 32(3):569-575, (1989) Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argeted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al., "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	0.01		Hynes et al., "further studies on the synthesis of quinazoline from 2-fluorobenzonitriles", J. Heterocyclic Chem.							
Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide argetted against C-raf Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	150		28:1357-1363, (1991)							
Kinase", Nature medicine, 2:668-675 (1996) Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal Oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	BC		Marsham et al., "Q	uinazoline Antifolate	Thymidylate Synthase Inh	ibitors", J. Me	d. Ckem. 32(3):569-575	5, (1989)	
Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal Oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	0.0	Monia et al., "antitumor activity of a phosphorothioate antisense oligedeoxynucleotide targeted against C-raf						C-raf		
Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Erec. Natl. Acad. Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal Oregulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			Kinase", Nature me	edicine, 2:668-675 (1	996)		3 4			
Sci. USA, 85:8855-8859 (1988) Robbins et al., "Regulation and properties of extracellular signal@regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			Morrison et al, "Signal transduction from membrane to cytolasm: Growth factors and membrane-bound							
Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro" J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	Pr		Oncogene products increase Raf-1 phophorylation and associated protein kinase activity", Proc. Natl. Acad.						\cad.	
J. Biol. Chem., 268:5097-5106, (1993) Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)			Sci. USA, 85:8855-8859 (19 8 8)							
Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	001		Robbins et al., "Regulation and properties of extracellular signal0regulated protein kinases 1 and 2 in vitro"							
Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)	150		J. Biol. Chem., 268:5097-5106, (1993)							
- Biological evaluation of a flevel class of o substitutes quintazonnelles , c. med. elistin, es(e). ee 1 te, (1886)	Webber et al., "Design of thymidylate synthase inhibitors using protein crystal structures: the synthesis and					and				
EVAMINED DATE CONCIDEDED	. 150	Biological evaluation of a novel class of 5-substituted quinazolinones", J. Med. Chem., 36(6):733-746, (1993)						(1993)		
	EXAMINER	<u> </u>	. 63		DATE CONSIDER		2002			
Brenda Coleman August 16, 2002									N. D.:	
 EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include any copy of this form with nex communication to applicant. 	line '	through	n citation if not in							